### PATENT COOPERATION TREATY

From the INTERNATIONAL SEARCHING AUTHORITY	PCT		
To: HANNA, MOORE & CURLEY Attn. Moore, Barry 11 Mespil Road Dublin 4 IRELAND - 4 NCV 2005	NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT AND THE WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY, OR THE DECLARATION		
Land a second of the second of	(PCT Rule 44.1)		
	Date of mailing (day/month/year) 04/11/2005		
Applicant's or agent's file reference M10-185-17PC	FOR FURTHER ACTION See paragraphs 1 and 4 below		
International application No. PCT/EP2005/050132	International filing date (day/month/year) 13/01/2005		
Applicant MICROSAIC SYSTEMS LIMITED			
1.   The applicant is hereby notified that the international search report and the written opinion of the International Searching Authority have been established and are transmitted herewith.  Filling of amendments and statement under Article 19: The applicant is entitled, if he so wishes, to amend the claims of the International Application (see Rule 46):  When? The time limit for filling such amendments is normally 2 months from the date of transmittat of the International Bureau of WIPO, 34 chemin des Colombettes  1211 Geneva 20, Switzerland, Fascimile No.: (41–22) 740.14.35  For more detailed instructions, see the notes on the accompanying sheet.  2. The applicant is hereby notified that no international search report will be established and that the dectaration under Article 17(2)(a) to that effect and the written opinion of the International Searching Authority are transmitted herewith.  3. With regard to the protest against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that:  the protest together with the decision thereon has been transmitted to the International Bureau together with the applicant's request to forward the texts of both the protest and the decision thereon to the designated Offices.  Reminders  Shortly after the expiration of 18 months from the protest; the applicant will be notified as soon as a decision is made.  4. Reminders  Shortly after the expiration of 18 months from the protest; the applicant will be notified as soon as a decision of the international gureau of the international gur			
Name and mailing address of the International Searching Authority  European Patent Office, P.B. 5818 Patentlaan 2  NL-2280 HV Rijswijk  Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  Fax: (431-70) 340-3016	Authorized officer Gennaro Cappiello		

#### NOTES TO FORM PCT/ISA/220

These Notes are intended to give the basic instructions concerning the filing of amendments under article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article", "Rule", and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions respectively.

#### INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international politication. Furthermore, it should be emphasized that provisional protection is available in some States only.

### What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

### When?

Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later, it should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Rule 46.1).

### Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been is filed, see below.

### How?

Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

### What documents must/may accompany the amendments?

Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

Notes to Form PCT/ISA/220 (first sheet) (January 1994)

### PATENT COOPERATION TREATY

## **PCT**

### INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference	FOR FURTHER	see Form PCT/ISA/220	
M10-185-17PC	ACTION	as well as, where applicable, item 5 below.	
International application No.	International filing date (day/mont)	/year) (Earliest) Priority Date (day/month/year)	
PCT/EP2005/050132	13/01/2005	12/02/2004	
Applicant			
MICROSAIC SYSTEMS LIMITED			
This International Search Report has been according to Article 18. A copy is being tra		rching Authority and is transmitted to the applicant	
This international Search Report consists	of a total ofsh	eets,	
X It is also accompanied by	a copy of each prior art document of	ited in this report.	
1. Basis of the report			
	international search was carried out ess otherwise indicated under this i	on the basis of the international application in the em.	
The international this Authority (Rul		of a translation of the international application furnished to	
b. With regard to any nucleo	otide and/or amino acid sequence	disclosed in the international application, see Box No. I.	
2. Certain claims were four	Certain claims were found unsearchable (See Box II).		
3. Unity of invention is lace	Unity of invention is lacking (see Box III).		
4. With regard to the title,			
X the text is approved as su	bmitted by the applicant.		
the text has been establis	hed by this Authority to read as folk	ws:	
į			
}			
1			
		1	
5. With regard to the abstract,			
the text is approved as su	•		
the text has been establis may, within one month from	thed, according to Rule 38.2(b), by the shall the date of mailing of this internation.	his Authority as it appears in Box No. IV. The applicant tional search report, submit comments to this Authority.	
6. With regard to the <b>drawings</b> ,			
a. the figure of the drawings to be p	oublished with the abstract is Figure	No	
X as suggested by			
1 =	s Authority, because the applicant f		
	is Authority, because this figure bett e published with the abstract	et Charactelitas die divendon.	
b. none of the figures is to b	e published with the abstract.		

### INTERNATIONAL SEARCH REPORT

International Application No PCT/EP2005/050132

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 H01J49/04				
.  According to International Patent Classification (IPC) or to both national classification and IPC				
B. FIELDS	SEARCHED			
Minimum do	cumentation searched (classification system followed by classification	n symbols)		
IPC 7	HOIJ	•	ļ	
Documentat	ion searched other than minimum documentation to the extent that se	uch documents are included in the fields se	arched	
Electronic d	ote has accounted during the international course (name of data books)			
	ata base consulted during the international search (name of data bas	e anu, where practical, search terms used)	•	
EPO-Internal, WPI Data				
	ENTS CONSIDERED TO BE RELEVANT			
Category °	Citation of document, with indication, where appropriate, of the rele	vant passages	Relevant to claim No.	
A	GB 2 026 231 A (EMI LTD) 30 January 1980 (1980-01-30) cited in the application the whole document		1-14	
A	LAPACK M A ET AL: "VALVED SAMPLING CELL 1 FOR MEMBRANE INTRODUCTION MASS SPECTROMETRY" ANALYTICAL CHEMISTRY, AMERICAN CHEMICAL SOCIETY. COLUMBUS, US,		1	
	vol. 68, no. 17, 1 September 1996 (1996-09-01), pa 3072-3075, XP000628535 ISSN: 0003-2700 the whole document	ges		
Α	GB 1 224 915 A (PADRTA) 10 March 1971 (1971-03-10) figure 1		1	
Furth	ner documents are listed in the continuation of box C.	X Patent family members are listed in	n annex.	
° Special ca	tegories of cited documents:	TI later degument published effectly inte	mational filling data	
"T' later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention				
"E" earlier document but published on or after the international filing date  "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone				
which is cited to establish the publication date of another citation or other special reason (as specified)  "O" document referring to an oral disclosure, use, exhibition or cannot be considered to involve an inventive step when the document referring to an oral disclosure, use, exhibition or				
other means ments, such combination being obvious to a person skilled in the art.				
tater than the priority date claimed "&" document member of the same patent ramity  Date of the actual completion of the international search  Date of mailing of the international search report				
19 October 2005 04/11/2005				
Name and mailing address of the ISA  Authorized officer  European Patent Office, P.B. 5818 Patentlaan 2				
NL - 2280 HV Rijswijk		Peters, V		

3 .

### INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No
PCT/EP2005/050132

Patent document cited in search report	Ī	Publication date		Patent family member(s)	Publication date
GB 2026231	Α	30-01-1980	NONE		
GB 1224915	A	10-03-1971	NONE		

### PATENT COOPERATION TREATY

From the INTERNATIONAL SEARCHING AUTHORITY To: WRITTEN OPINION OF THE see form PCT/ISA/220 INTERNATIONAL SEARCHING AUTHORITY (PCT Rule 43bis.1) Date of mailing (day/month/year) see form PCT/ISA/210 (second sheet) Applicant's or agent's file reference FOR FURTHER ACTION see form PCT/ISA/220 See paragraph 2 below International application No. International filing date (day/month/year) Priority date (day/month/year) PCT/EP2005/050132 12.02.2004 13.01.2005 International Patent Classification (IPC) or both national classification and IPC H01J49/04 Applicant MICROSAIC SYSTEMS LIMITED This opinion contains indications relating to the following items: Box No. I Basis of the opinion □ Box No. II Priority Box No. III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability Box No. IV Box No. V Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement ☐ Box No. VI Certain documents cited ☐ Box No. VII Certain defects in the international application ☑ Box No. VIII Certain observations on the international application **FURTHER ACTION** If a demand for international preliminary examination is made, this opinion will usually be considered to be a written opinion of the International Preliminary Examining Authority ("IPEA"). However, this does not apply where the applicant chooses an Authority other than this one to be the IPEA and the chosen IPEA has notifed the International Bureau under Rule 66.1 bis(b) that written opinions of this International Searching Authority will not be so considered. If this opinion is, as provided above, considered to be a written opinion of the IPEA, the applicant is invited to submit to the IPEA a written reply together, where appropriate, with amendments, before the expiration of three months from the date of mailing of Form PCT/ISA/220 or before the expiration of 22 months from the priority date, whichever expires later. For further options, see Form PCT/ISA/220. For further details, see notes to Form PCT/ISA/220. Name and mailing address of the ISA: Authorized Officer

Peters, V

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NL-2280 HV Rijswijk - Pays Bas Tel. +31 70 340 - 2040 Tx: 31 651 epo nl

Fax: +31 70 340 - 3016

# WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No. PCT/EP2005/050132

_	Box N	o. ! Basis of the opinion
1.		egard to the <b>language</b> , this opinion has been established on the basis of the international application in guage in which it was filed, unless otherwise indicated under this item.
	lai	nis opinion has been established on the basis of a translation from the original language into the following inguage—, which is the language of a translation furnished for the purposes of international search inder Rules 12.3 and 23.1(b)).
2.		egard to any <b>nucleotide and/or amino acid sequence</b> disclosed in the international application and sary to the claimed invention, this opinion has been established on the basis of:
	a. type	of material:
		a sequence listing
		table(s) related to the sequence listing
	b. form	nat of material:
		in written format
		in computer readable form
	c. time	of filing/furnishing:
		contained in the international application as filed.
		filed together with the international application in computer readable form.
		furnished subsequently to this Authority for the purposes of search.
3.	ha Co	addition, in the case that more than one version or copy of a sequence listing and/or table relating thereto as been filed or furnished, the required statements that the information in the subsequent or additional opies is identical to that in the application as filed or does not go beyond the application as filed, as oppopriate, were furnished.
4.	Addition	onal comments:

# WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No. PCT/EP2005/050132

Box No. III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability				
The questions whether the claimed invention appears to be novel, to involve an inventive step (to be non obvious), or to be industrially applicable have not been examined in respect of:				
☐ the entire international application	the entire international application,			
⊠ claims Nos. 14				
because:				
the said international application, or the said claims Nos. relate to the following subject matter which does not require an international preliminary examination (specify):				
	the description, claims or drawings (indicate particular elements below) or said claims Nos. are so unclear that no meaningful opinion could be formed (specify):			
the claims, or said claims Nos. are could be formed.				
□ no international search report has	no international search report has been established for the whole application or for said claims Nos. 14			
	the nucleotide and/or amino acid sequence listing does not comply with the standard provided for in Annex C of the Administrative Instructions in that:			
the written form	has not been furnished			
	does not comply with the standard			
the computer readable form	has not been furnished			
	does not comply with the standard			
	e and/or amino acid sequence listing, if in computer readable form only, do sirements provided for in Annex C-bis of the Administrative Instructions.			
☐ See separate sheet for further det	rails			

# WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No. PCT/EP2005/050132

Box No. V Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Statement

Novelty (N)

Yes: Claims

1-13

No: Claims

Inventive step (IS)

Yes: Claims

Claims

1-13

Industrial applicability (IA)

Yes: Claims

1-14

No: Claims

No:

2. Citations and explanations

see separate sheet

Box No. VIII Certain observations on the international application

The following observations on the clarity of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made:

see separate sheet

### Re Item III.

Since it is not clear for which features of the description and drawings protection is sought, no meaningful search of claim 14 is possible (see item VIII of the present opinion).

### Re Item V.

1 Reference is made to the following documents:

D1: GB 2 026 231 A (EMI LTD) 30 January 1980 (1980-01-30)

D2: LAPACK M A ET AL: "VALVED SAMPLING CELL FOR MEMBRANE INTRODUCTION MASS SPECTROMETRY" ANALYTICAL CHEMISTRY, AMERICAN CHEMICAL SOCIETY. COLUMBUS, US, vol. 68, no. 17, 1 September 1996 (1996-09-01), pages 3072-3075, XP000628535 ISSN: 0003-2700

2 Document D1, which is considered to represent the most relevant state of the art, discloses (the references in parentheses applying to this document):

A mass spectrometer system (cl. 1) including a mass spectrometer device (cl. 1) provided within an evacuated chamber (cl. 1), the chamber having an entrance port (13) through which a sample may be introduced into the chamber and into contact with the mass spectrometer device, the system additionally including a permeable membrane (14) located across the port and a valve (12) located between the membrane and the spectrometer device and having a closed state and an open state, such that, in use, the adoption of the open state allows the flow of the sample into the chamber through the membrane and into contact with the spectrometer device.

From this, the subject-matter of independent claim 1 differs in that the permeable membrane is located across the chamber between the port and the spectrometer device, the valve is located between the entrance port and the membrane and the valve is normally in its closed state.

The subject-matter of claim 1 is therefore novel (Article 33(2) PCT)

2.1 The problem to be solved by the present invention may be regarded as providing

# WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY (SEPARATE SHEET)

International application No.

PCT/EP2005/050132

an alternative arrangement for removing the need to include a system to provide the vacuum necessary for operation of the mass spectrometer (see p. 2, I. 9-13 of the present application).

- 2.2 The solution to this problem proposed in claim 1 of the present application is considered as involving an inventive step (Article 33(3) PCT) because non of the documents in the prior art disclose or make obvious the solution presented in the current application.
- 2.3 Claims 2-13 are dependent on claim 1 and as such also meet the requirements of the PCT with respect to novelty and inventive step.

### Re Item VIII.

The claims shall not rely, in respect of the technical features of the invention, on references to the description or drawings (Rule 6.2(a) PCT; Guidelines 5.10).

### NOTES TO FORM PCT/ISA/220 (continued)

The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled:
- (iii) the claim is new;
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

### The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- [Where originally there were 48 claims and after amendment of some claims there are 51]:
   "Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers;
   claims 30, 33 and 36 unchanged; new claims 49 to 51 added."
- Where originally there were 15 claims and after amendment of all claims there are 11]: "Claims 1 to 15 replaced by amended claims 1 to 11."
- [Where originally there were 14 claims and the amendments consist in cancelling some claims and in adding new claims]:
   "Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added." or "Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged."
- [Where various kinds of amendments are made]:
   "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claim 14; claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

#### "Statement under article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on the description and the drawings (which cannot be amended under Article 19(1)).

The statement will be published with the international application and the amended claims.

It must be in the language in which the international appplication is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the letter indicating the differences between the claims as filed and as amended. It must be filed on a separate sheet and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

### Consequence if a demand for international preliminary examination has already been flied

If, at the time of filing any amendments under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the same time of filing the amendments with the International Bureau, also file a copy of such amendments with the International Preliminary Examining Authority (see Rule 62.2(a), first sentence).

### Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, where upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

For further details on the requirements of each designated/elected Office, see Volume II of the PCT Applicant's Guide

Notes to Form PCT/ISA/220 (second sheet) (January 1994)